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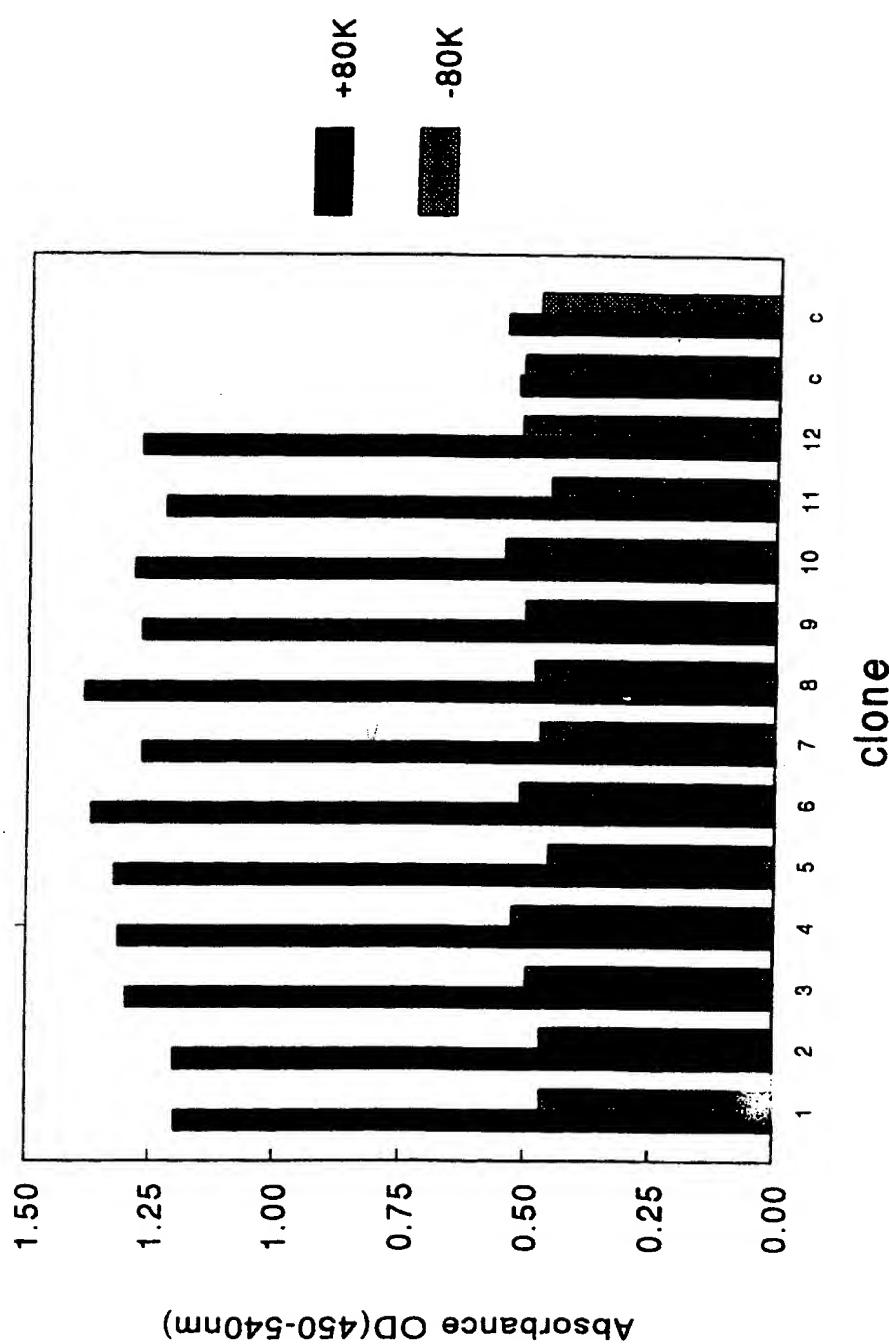


Figure 1

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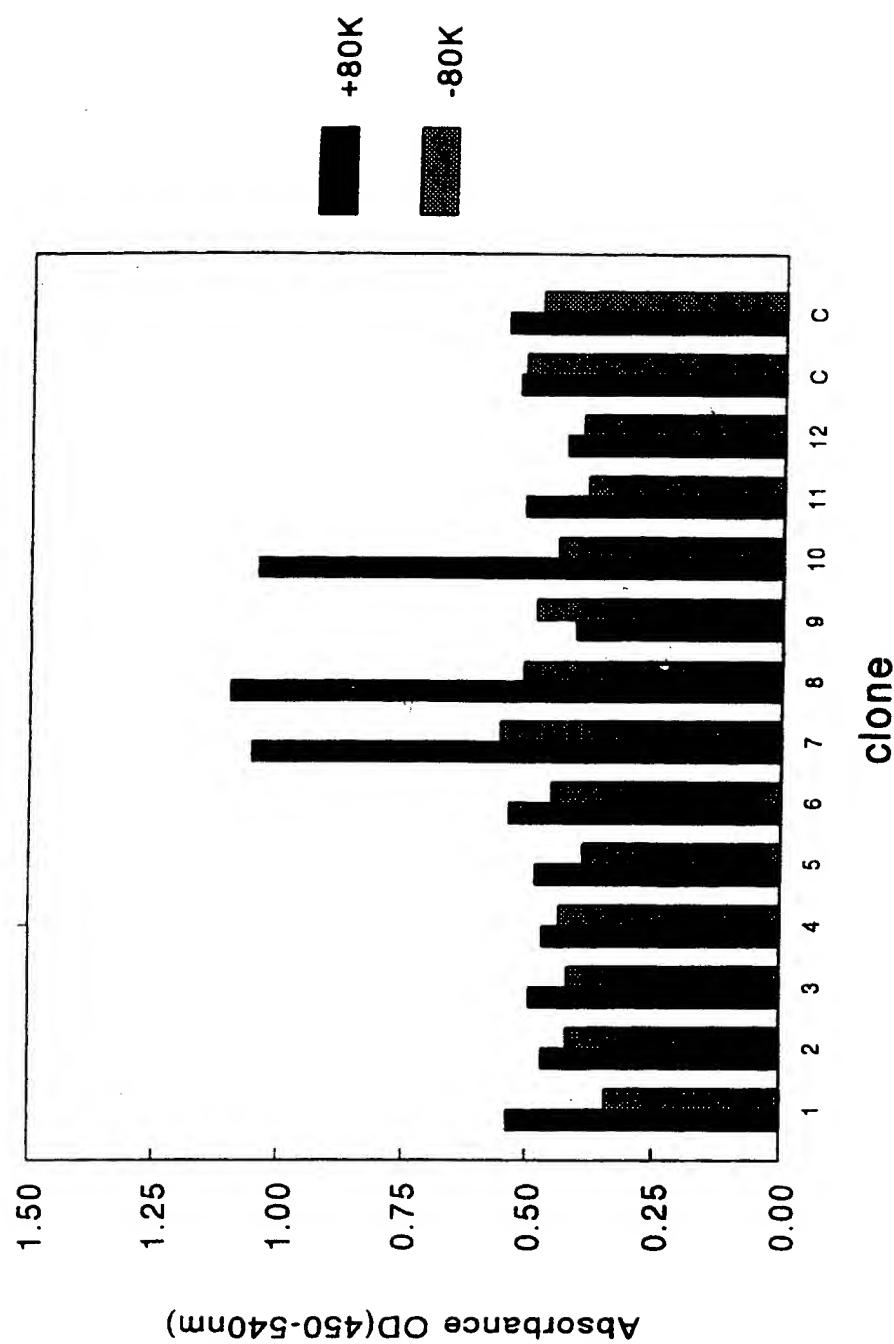


Figure 2

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	CDR1	CDR2	CDR3	CDR4	CDR1	CDR2	CDR3	CDR4	
FL-14	1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23 24 25 26 27 28 29 30 31 31a 31b 32 33 34 35 36 37 38 39 40 41 42 43 44 45 46 47 48 49 CAG GTC CAG CTG GTC CAG TGT GCG GTC AGG CCT CGG TCC TGG AGG GTC TGC AGG GAC ACC TTC AAC NOC ... ... TTT CCT ATC AGT TGG CCA CAG GCC CCT GAA CAA TAT TAC TGG ATG GGA 10	50 51 52 52a 52b 52c 53 54 55 56 57 58 59 60 61 62 63 64 65 66 67 68 69 70 71 72 73 74 75 76 77 78 79 80 81 82 82a 82b 82c 83 84 85 86 87 88 89 90 91 92 93 94 Gcc ATC ATC CCT ... ... ATC TTT GGT TCA ACA AGG TAC CGA CGG AGG TTC AGG GTC AGG CTC GAT GCT GCA AGA GTC AGC ACC TCG AAC ACC TCG AGC TAC AGA TCA TAT AAC TGT GCA CGA 10	95 96 97 98 99 100 a b c d e f g h i j k l m 101 102 103 104 105 106 107 108 109 110 111 112 113 CAA CAG AAC GGC AGC TGG TAC GAA GGA CGG TGG CTT GAG CGG AGG CCT GAT GCT GTC AGG GTC TCC TGA AGG GTC ACC GTC ACC GTC AGT G,	50 51 52 52a 52b 52c 53 54 55 56 57 58 59 60 61 62 63 64 65 66 67 68 69 70 71 72 73 74 75 76 77 78 79 80 81 82 82a 82b 82c 83 84 85 86 87 88 89 90 91 92 93 94 TGG ATC AGC ATT ... ... TAT AGT GGT AAC AGC TAC GCA AGG TCC AGG AGA GTC AGC ACC AGG AAC AGA TCC AGC AGA AAC GCA AGC TAC AGC TCG AGA TCT GAC AGC TAC TAT TGT GGC AGA 14	95 96 97 98 99 100 a b c d e f g h i j k l m n o 101 102 103 104 105 106 107 108 109 110 111 112 113 GAT GGG GGG GGG CCT GCC TAT GAA GAT GTT TGG AGT GGT GAG TAC CCC GAA TAC TAC GCT ATG GTC TGG GTC ACC AGC GTC ACC GTC AAC GCA				

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**Figure 4A**

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FR1	CDR1	FR2	CDR2	FR3	CDR3	FR4
123456789012345678901234567890	1ab2345	67890123456789	012abc3456789012345	67890123456789012345678901234	567890abcdefgijklmnno12	34567890123
DP-14 VH IT-2 VH EL-5 VH EL-25	QVQLVOSGAEVKKPGASVKVSCKASGYTFT ----L--AT----M----M----P---- ----L--AT----M----M----R---- ----L--A----	S--YGIS WVRQARGQGLEHMG ----D-- ----D--	WISAA--YNGNTNYAQKLOG RVYMTTDSTSAYMEIRSLSRSDDTAVYYCAR ----I---S----D----E---- ----I---S----D----F----	DGGGAYEDVMSGEYPEYYAMDY	WGQGTTVTVSS	
				----V---- ----D--	----RR---- ----RR----	

Figure 4B

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CDR3

EL-14      QQNQGGWYEGPLLEPRPD--ALDI  
              | | | | . | . | . | . | . |  
IT-2      DGGGGAYEDVWSGEYPEYYAMDV

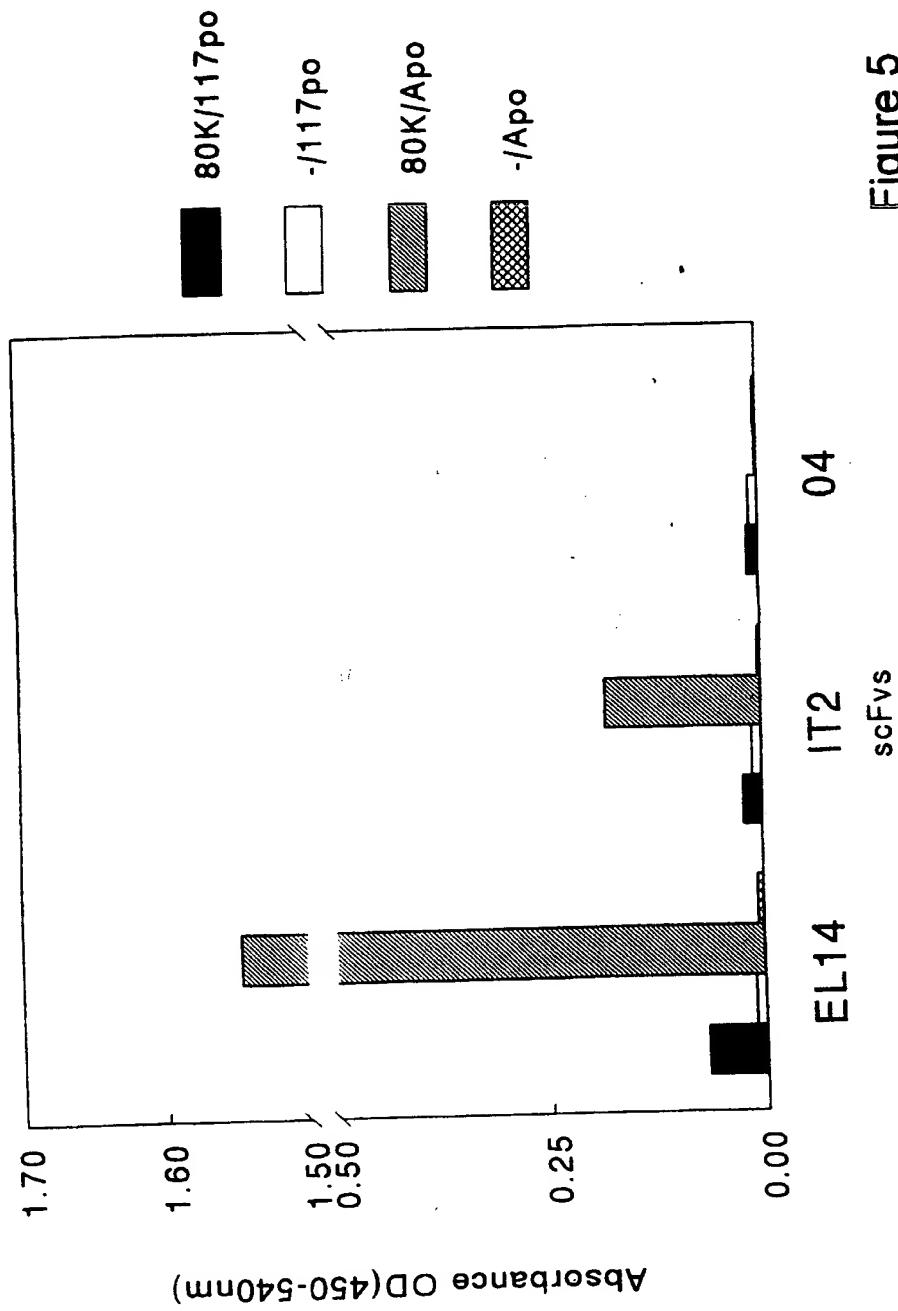
Figure 4C

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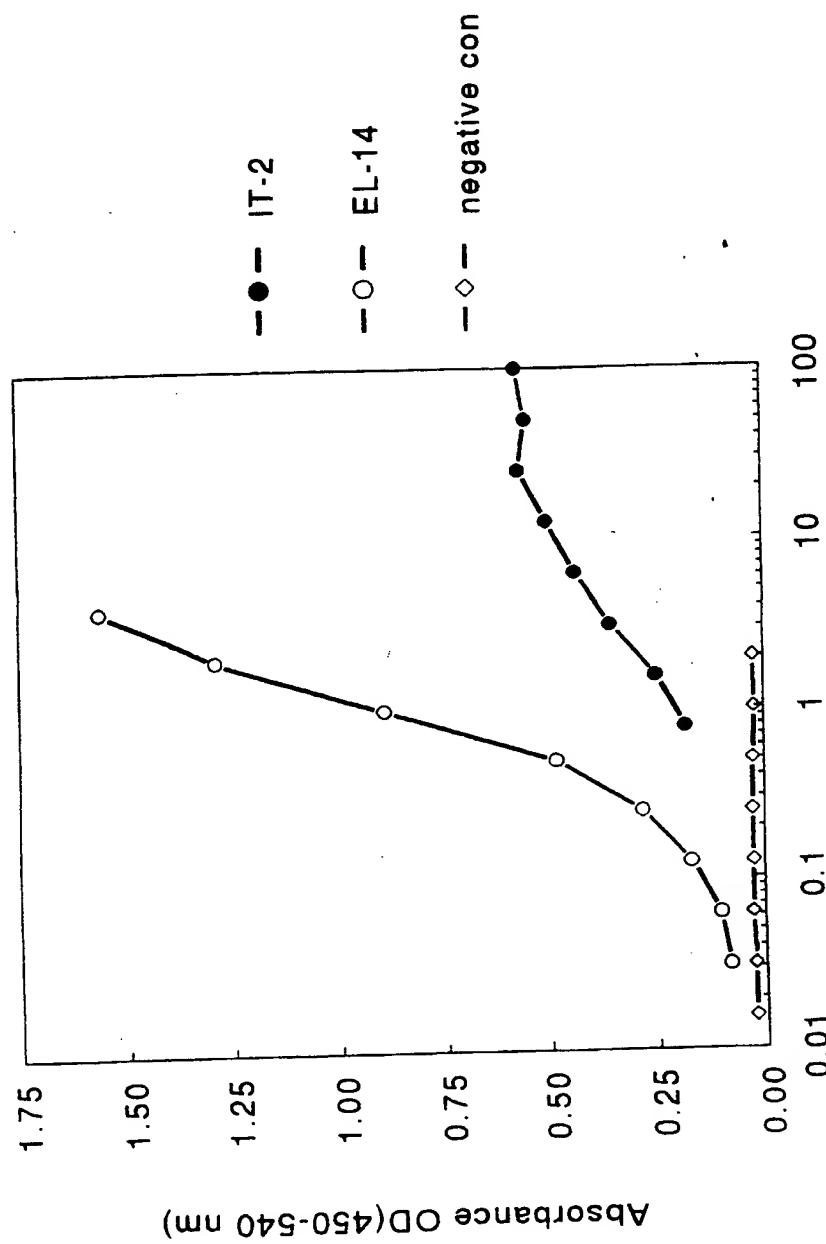


Figure 6

Concentration ( $\mu\text{g}/\text{ml}$ )

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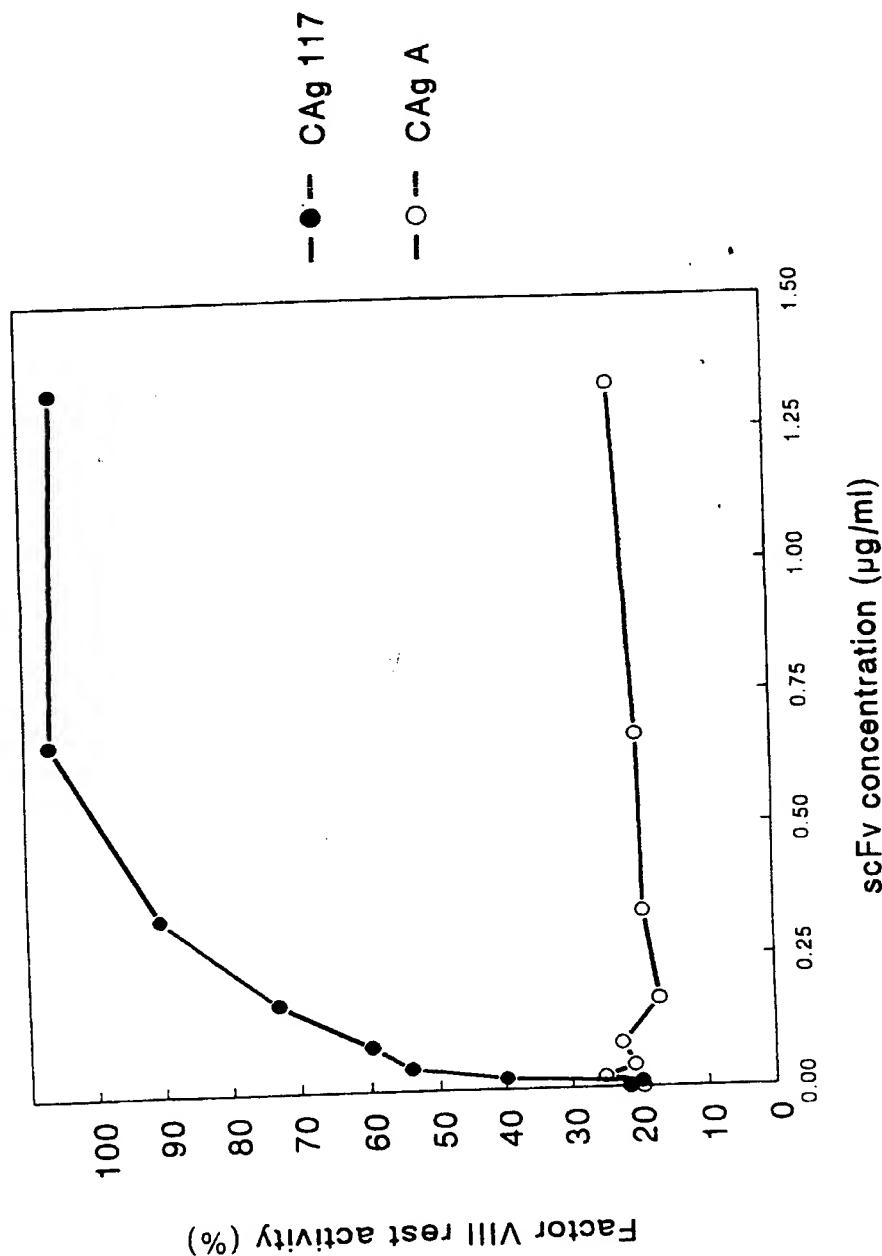


Figure 7A NEUTRALIZATION BY scFv EL-14

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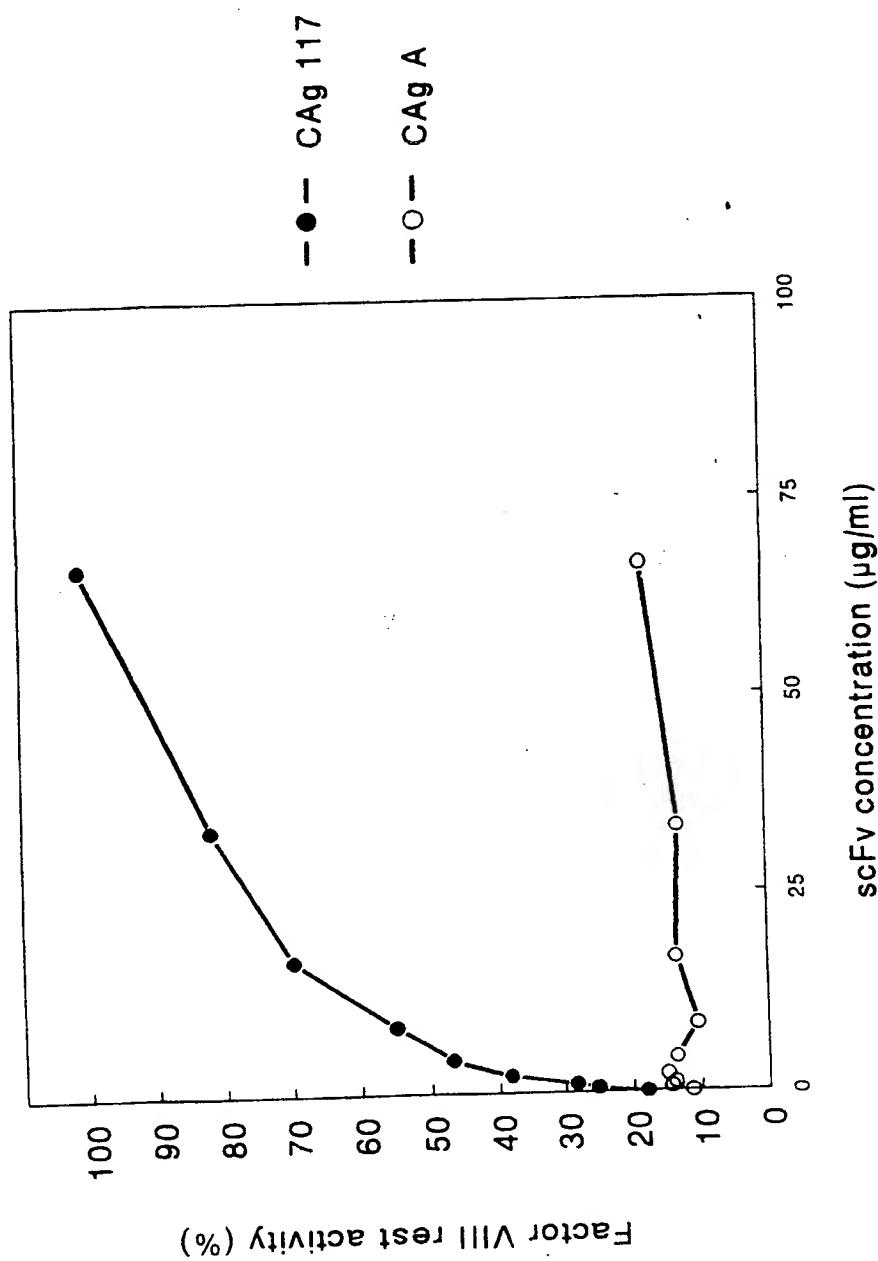
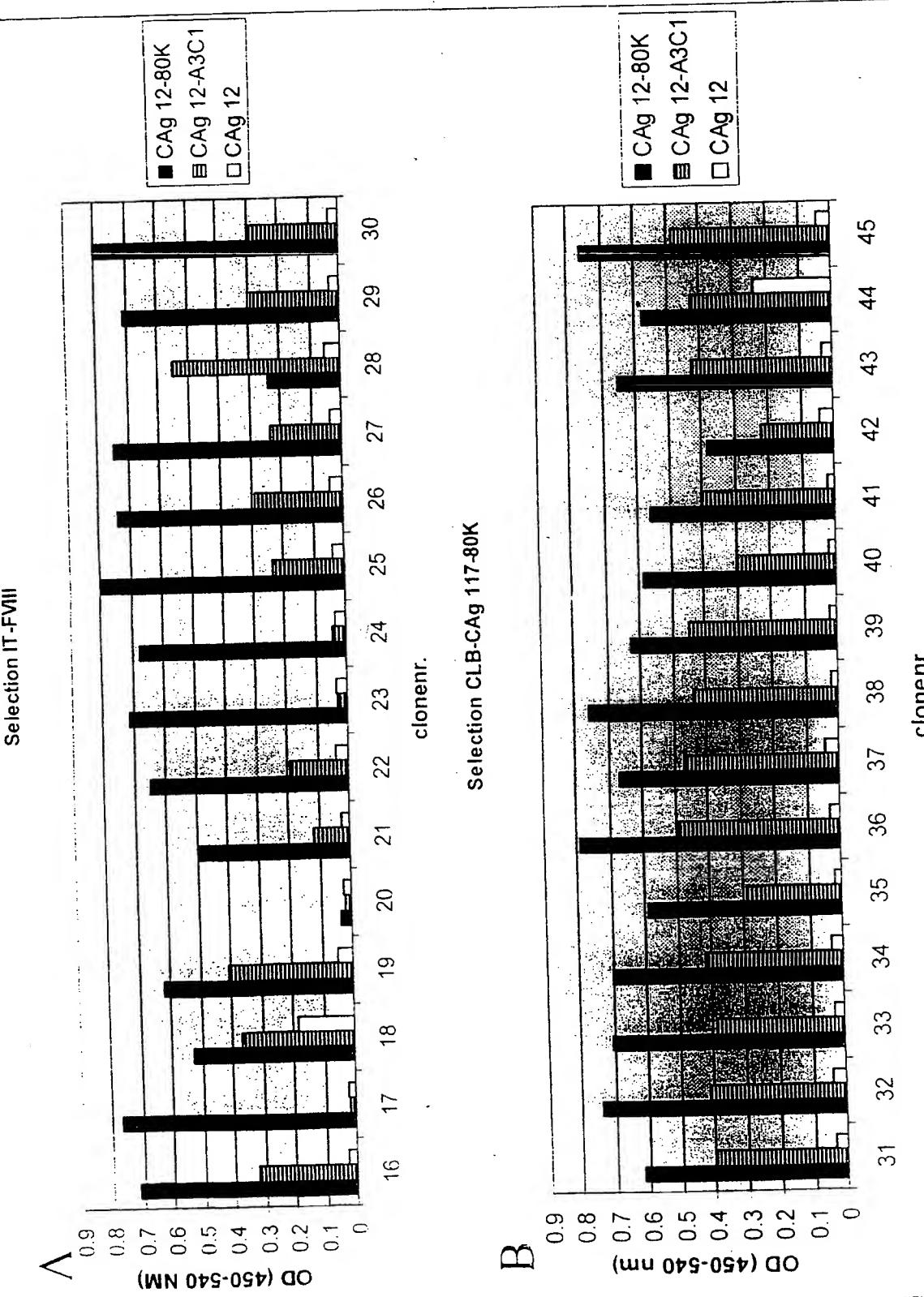


Figure 7B NEUTRALIZATION BY scFv IT-2

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Figure 8



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## Deduced protein sequences of isolated FVIII A3-C1 specific scFv

Heavy chains	FR1	CDR1	FR2	CDR2	FR3	CDR3	FR4
12345678901234567890 12345 678901234567890 012a3456789012345 67890123456789012abc345678901234 567890abdefghil2 34567890123	1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3				1 9 0 1 0 1 1		
DP-15 QVQLVQSGAEVKKEPGASVKSCKASSYIFT B38	-----L-V-A-D-----T-----I-----	-----A-F-----K-----	-----A-P-----K-----	-----A-----F-----	-----Y-----K-----	-----RN-----E-----	-----CDTLLLIWFGPAPNDS WGGTTLV
DP-31 EYOLVESGGGLVQPGRSLRLSCAASGFTFD B18	-----Q-----Q-----K-----G-----I-----E-----	-----V-----D-----L-----	-----E-----I-----A-----	-----F-----F-----A-----	-----Y-----A-----A-----	-----L-----A-----A-----	-----DPIINSSNYRRGVAAFDI WGGQTMTVSS
DP-49 QVQLVESGGGVVQPGRSLRLSCAASGFTFS B35	-----E-----L-----K-----V-----D-----L-----	-----A-----A-----A-----	-----A-----A-----A-----	-----ND-----ND-----	-----A-----A-----A-----	-----TI-----TI-----	-----DLIESNI AEAL WGGQTTLVTSS
DP-77 EVQLVESGGGLVKPQGSPLRLSCAASGFTFS B04	-----R-----K-----E-----	-----R-----R-----	-----D-----T-----	-----T-----G-----G-----	-----M-----N-----V-----	-----F-----F-----	-----DGTFGSAATWR AFDI WGRGTMTVSSG

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+1    -Gln<sub>1</sub> -Val<sub>2</sub> -Gln<sub>3</sub> -Leu<sub>4</sub> -Leu<sub>5</sub> -Gln<sub>6</sub> -Ser<sub>7</sub> -Ala<sub>8</sub> -Ala<sub>9</sub> -Asp<sub>10</sub> -Val<sub>11</sub> -Lys<sub>12</sub> -Lys<sub>13</sub> -Pro<sub>14</sub> -Gly<sub>15</sub> -Ala<sub>16</sub> -Ser<sub>17</sub>  
 CAGGTGCAGC TGTTGCAGTC TGCAGCTGAC GTGAAGAACG CTGGGGCCTC 50  
 GTCCACGTAG ACAACGTCAG ACGTCGACTG CACCTTCTTCG GACCCCGGAG

+1    - -Val<sub>1</sub> -Lys<sub>2</sub> -Val<sub>3</sub> -Ser<sub>4</sub> -Cys<sub>5</sub> -Thr<sub>6</sub> -Ala<sub>7</sub> -Ser<sub>8</sub> -Gly<sub>9</sub> -Tyr<sub>10</sub> -Ile<sub>11</sub> -Phe<sub>12</sub> -Thr<sub>13</sub> -Ser<sub>14</sub> -Tyr<sub>15</sub> -Asp<sub>16</sub> -Ile<sub>17</sub>  
 AGTGAAGGTC TCCTGTACGG CTTCTGGATA CATCTTCACC AGTTATGATA 100  
 TCACCTCCAG AGGACATGCC GAAGACCTAT GTAGAAGTGG TCAATACTAT

+1    - -Asn<sub>1</sub> -Trp<sub>2</sub> -Val<sub>3</sub> -Arg<sub>4</sub> -Gln<sub>5</sub> -Ala<sub>6</sub> -Thr<sub>7</sub> -Gly<sub>8</sub> -Gly<sub>9</sub> -Leu<sub>10</sub> -Glu<sub>11</sub> -Trp<sub>12</sub> -Met<sub>13</sub> -Gly<sub>14</sub> -Trp<sub>15</sub>  
 TCAAACGGGT GCGACAGGCC ACTGGACAAG GGCTTGAGTG GATGGGATGG 150  
 AGTTGACCCA CGCTGTCCGG TGACCTGTTG CCGAACTCAC CTACCCCTACC

+1    - -Met<sub>1</sub> -Asn<sub>2</sub> -Pro<sub>3</sub> -Asn<sub>4</sub> -Ser<sub>5</sub> -Gly<sub>6</sub> -Asp<sub>7</sub> -Ala<sub>8</sub> -Gly<sub>9</sub> -Phe<sub>10</sub> -Ala<sub>11</sub> -Gln<sub>12</sub> -Lys<sub>13</sub> -Phe<sub>14</sub> -Lys<sub>15</sub> -Gly<sub>16</sub> -Arg<sub>17</sub>  
 ATGAATCCTA ACAGTGGTAA CGCAGGCTTT GCACAGAAAG TTAAGGGCAG 200  
 TACTTAGGAT TGTCACCATT GCGTCCGAAA CGTGTCTTC AATTCCCGTC

+1    - -Leu<sub>1</sub> -Thr<sub>2</sub> -Leu<sub>3</sub> -Thr<sub>4</sub> -Arg<sub>5</sub> -Asp<sub>6</sub> -Thr<sub>7</sub> -Ser<sub>8</sub> -Thr<sub>9</sub> -Ala<sub>10</sub> -Tyr<sub>11</sub> -Met<sub>12</sub> -Glu<sub>13</sub> -Leu<sub>14</sub> -Arg<sub>15</sub>  
 ACTCACCTTG ACCAGGGACA CTTCCACAAG CACAGCCTAC ATGGAGCTGA 250  
 TGAGTGGAAC TGGTCCCTGT GAAGGTGTTG GTGTCGGATG TACCTCGACT

+1    - -Arg<sub>1</sub> -Leu<sub>2</sub> -Glu<sub>3</sub> -Ser<sub>4</sub> -Glu<sub>5</sub> -Asp<sub>6</sub> -Thr<sub>7</sub> -Ala<sub>8</sub> -Val<sub>9</sub> -Tyr<sub>10</sub> -Tyr<sub>11</sub> -Cys<sub>12</sub> -Ala<sub>13</sub> -Arg<sub>14</sub> -Cys<sub>15</sub> -Asp<sub>16</sub>  
 GGAGACTGGA ATCTGAGGAC ACGGCCGTGT ATTACTGTGC GAGATGTGAC 300  
 CCTCTGACCT TAGACTCCTG TGCCGGCACA TAATGACACG CTCTACACTG

+1    - -Thr<sub>1</sub> -Thr<sub>2</sub> -Leu<sub>3</sub> -Ile<sub>4</sub> -Trp<sub>5</sub> -Phe<sub>6</sub> -Gly<sub>7</sub> -Pro<sub>8</sub> -Ala<sub>9</sub> -Pro<sub>10</sub> -Tyr<sub>11</sub> -Tyr<sub>12</sub> -Asp<sub>13</sub> -Ser<sub>14</sub> -Trp<sub>15</sub> -Gly<sub>16</sub>  
 ACCACACTCT TAATCTGGTT CGGGCCCCGCC CCCTACTATG ACTCCTGGGG 350  
 TGGTGTGAGA ATTAGACCAA GCCCGGGCGG GGGATGATAC TGAGGACCCC

+1    - -Gln<sub>1</sub> -Gly<sub>2</sub> -Thr<sub>3</sub> -Leu<sub>4</sub> -Val<sub>5</sub>  
 CCAGGGAACCTAGTC  
 GGTCCCTTGA GATCAG

400

Figure 9B

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+1    rGln, rVal, rGln, rLeu, rVal, rGln, rSer, rGly, rGly, rGly, rLeu, rVal, rGln, rPro, rGly, rLys, rSer-  
 CAGGTGCAAC TGGTGCAGTC TGGGGGAGGC TTGGTACAGC CTGGCAAGTC 50  
 GTCCACGTTG ACCACGTCAG ACCCCCCTCCG AACCATGTCG GACCGTTAG

+1    rLeu, rArg, rLeu, rSer, rCys, rAla, rAla, rSer, rGly, rPhe, rThr, rPhe, rGly, rAsp, rTyr, rAla, rIle-  
 CCTGAGACTC TCCTGTGCAG CCTCTGGATT CACATTTGGC GATTATGCCA 100  
 GGACTCTGAG AGGACACGTC GGAGACCTAA GTGTAAAACCG CTAATACGGT

+1    —, rHis, rTrp, rVal, rArg, rGln, rAla, rPro, rGly, rGlu, rGly, rLeu, rGlu, rTrp, rVal, rSer, rGly-  
 TACACTGGGT CCGGCAAGCT CCAGGGGAGG GCCTGGAGTG GGTCTCAGGT 150  
 ATGTGACCCA GGCGTTCGA GGTCCCCCTCC CGGACCTCAC CCAGAGTCCA

+1    rVal, rThr, rTrp, rSer, rGly, rThr, rIle, rGly, rPhe, rAla, rAsp, rSer, rVal, rLys, rGly, rArg-  
 GTTACTTGGGA GTGGTACTAC TATAGGCTTT GCGGACTCTG TGAAGGGCCG 200  
 CAATGAACCT CACCATGATG ATATCCGAAA CGCCTGAGAC ACTTCCCAGC

+1    rPhe, rThr, rIle, rSer, rArg, rAsp, rAsn, rAla, rLys, rAsn, rSer, rLeu, rTyr, rLeu, rTyr, rMet, rAsn-  
 ATTCAACCATC TCCAGAGACA ACGCCAAGAA TTCCCTGTAT CTGTACATGA 250  
 TAAGTGGTAG AGGTCTCTGT TGCGGTTCTT AAGGGACATA GACATGTACT

+1    —, rSer, rLeu, rArg, rAla, rGlu, rAsp, rThr, rAla, rLeu, rTyr, rTyr, rCys, rAla, rLeu, rPro, rTyr-  
 ACAGTCTGAG AGCTGAAGAC ACGGCCTTGT ATTATTGTGC CTTACCATAT 300  
 TGTCAGACTC TCGACTTCTG TGCCGGAACA TAATAACACG GAATGGTATA

+1    —, rIle, rAsn, rSer, rSer, rAsn, rTyr, rArg, rArg, rGly, rVal, rAla, rAla, rPhe, rAsp, rIle, rTrp, rGly-  
 ATCAAACCTCGT CCAAACCTACAG AAGAGGGGTC GCTGCCTTCG ATATCTGGGG 350  
 TAGTTGAGCA GGTTGATGTC TTCTCCCCAG CGACGGAAAGC TATAGACCCC

+1    —, rGln, rGly, rThr, rMet, rVal, rThr, rVal, rSer, rSer  
 CCAAGGGACA ATGGTCACCG TGTCGAGT 400  
 GGTTCCCTGT TACCAAGTGGC ACAGCTCA

Figure 9C

+1 - Glu<sub>1</sub> - Val<sub>2</sub> - Glu<sub>3</sub> - Leu<sub>4</sub> - Val<sub>5</sub> - Glu<sub>6</sub> - Ser<sub>7</sub> - Gly<sub>8</sub> - Gly<sub>9</sub> - Gly<sub>10</sub> - Leu<sub>11</sub> - Val<sub>12</sub> - Gln<sub>13</sub> - Pro<sub>14</sub> - Gly<sub>15</sub> - Arg<sub>16</sub> - Ser<sub>17</sub>  
 GAGGTGCAGC TGGTGGAGTC TGGGGGAGGC TTGGTACAGC CTGGGAGGTC 50  
 CTCCACGTCG ACCACCTCAG ACCCCCCTCCG AACCATGTCA GACCCTCCAG

+1 - Leu<sub>1</sub> - Arg<sub>2</sub> - Leu<sub>3</sub> - Ser<sub>4</sub> - Cys<sub>5</sub> - Val<sub>6</sub> - Asp<sub>7</sub> - Ser<sub>8</sub> - Gly<sub>9</sub> - Leu<sub>10</sub> - Thr<sub>11</sub> - Phe<sub>12</sub> - Ser<sub>13</sub> - Ser<sub>14</sub> - Tyr<sub>15</sub> - Gly<sub>16</sub> - Met<sub>17</sub>  
 CCTGAGACTC TCCTGTGTAG ACTCTGGACT CACCTTCAGT AGTTATGGCA 100  
 GGACTCTGAG AGGACACATC TGAGACCTGA GTGGAAGTCA TCAATACCGT

+1 - His<sub>1</sub> - Trp<sub>2</sub> - Val<sub>3</sub> - Arg<sub>4</sub> - Gln<sub>5</sub> - Ala<sub>6</sub> - Pro<sub>7</sub> - Gly<sub>8</sub> - Ala<sub>9</sub> - Gly<sub>10</sub> - Leu<sub>11</sub> - Glu<sub>12</sub> - Trp<sub>13</sub> - Val<sub>14</sub> - Ala<sub>15</sub> - Val<sub>16</sub>  
 TGCACTGGGT CCGCCAGGCT CCAGGCCGCG GGCTGGAGTG GGTGGCCGTT 150  
 ACGTGACCCA GGCAGTCCGA GGTCCCGCCT CCGACCTCAC CCACCGGCAA

+1 - Ile<sub>1</sub> - Ser<sub>2</sub> - Tyr<sub>3</sub> - Asp<sub>4</sub> - Gly<sub>5</sub> - Asn<sub>6</sub> - Asp<sub>7</sub> - Lys<sub>8</sub> - Tyr<sub>9</sub> - Ala<sub>10</sub> - Asp<sub>11</sub> - Ser<sub>12</sub> - Val<sub>13</sub> - Lys<sub>14</sub> - Gly<sub>15</sub> - Arg<sub>16</sub>  
 ATTCATACG ACGGAAATGA TAAATATTAT GCAGACTCCG TGAAGGGCCG 200  
 TAAAGTATGC TGCCTTACT ATTTATAATA CGTCTGAGGC ACTTCCCCGC

+1 - Phe<sub>1</sub> - Ala<sub>2</sub> - Ile<sub>3</sub> - Ser<sub>4</sub> - Arg<sub>5</sub> - Asp<sub>6</sub> - Asn<sub>7</sub> - Ala<sub>8</sub> - Lys<sub>9</sub> - Asn<sub>10</sub> - Thr<sub>11</sub> - Leu<sub>12</sub> - Tyr<sub>13</sub> - Leu<sub>14</sub> - Gln<sub>15</sub> - Met<sub>16</sub> - Asn<sub>17</sub>  
 ATTGCCCATC TCCAGAGACA ATGCCAAGAA CACGCTGTAT CTGCAAATGA 250  
 TAAGCGGTAG AGGTCTCTGT TACGGTTCTT GTGCGACATA GACGTTTACT

+1 - Ser<sub>1</sub> - Leu<sub>2</sub> - Thr<sub>3</sub> - Ile<sub>4</sub> - Glu<sub>5</sub> - Asp<sub>6</sub> - Thr<sub>7</sub> - Ala<sub>8</sub> - Val<sub>9</sub> - Tyr<sub>10</sub> - Cys<sub>11</sub> - Ala<sub>12</sub> - Lys<sub>13</sub> - Asp<sub>14</sub> - Leu<sub>15</sub>  
 ACAGCCTGAC AATAGAGGAC ACGGCTGTCT ATTATTGTGC GAAAGATCTC 300  
 TGTCGGACTG TTATCTCCTG TGCCGACAGA TAATAACACG CTTCTAGAG

+1 - Ile<sub>1</sub> - Glu<sub>2</sub> - Ser<sub>3</sub> - Asn<sub>4</sub> - Ile<sub>5</sub> - Ala<sub>6</sub> - Glu<sub>7</sub> - Ala<sub>8</sub> - Leu<sub>9</sub> - Thr<sub>10</sub> - Gly<sub>11</sub> - Gln<sub>12</sub> - Gly<sub>13</sub> - Thr<sub>14</sub> - Leu<sub>15</sub> - Val<sub>16</sub> - Thr<sub>17</sub>  
 ATAGAACCAA ATATTGCGGA GGCCCTCTGG GGCCAGGGAA CCCTGGTCAC 350  
 TATCTTAGTT TATAACGCCT CCGGGAGACC CCGGTCCCTT GGGACCAGTG

+1 - Val<sub>1</sub> - Ser<sub>2</sub> - Ser<sub>3</sub>  
 CGTGTGAGT  
 GCACAGCTCA

400

Figure 9D

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+1 : Glu<sub>1</sub> Val<sub>2</sub> Gln<sub>3</sub> Leu<sub>4</sub> Val<sub>5</sub> Lys<sub>6</sub> Ser<sub>7</sub> Gly<sub>8</sub> Glu<sub>9</sub> Gly<sub>10</sub> Leu<sub>11</sub> Val<sub>12</sub> Lys<sub>13</sub> Pro<sub>14</sub> Gly<sub>15</sub> Gly<sub>16</sub> Ser-  
GAGGTGCAGC TGGTGAAGTC TGGGGAAGGC CTGGTCAAGC CTGGGGGGTC 50  
CTCCACGTCG ACCACTTCAG ACCCCTCCG GACCAGTCG GACCCCCCAG

-1 : - Leu<sub>1</sub> Arg<sub>2</sub> Leu<sub>3</sub> Ser<sub>4</sub> Cys<sub>5</sub> Ala<sub>6</sub> Ala<sub>7</sub> Ser<sub>8</sub> Gly<sub>9</sub> Phe<sub>10</sub> Thr<sub>11</sub> Phe<sub>12</sub> Arg<sub>13</sub> Arg<sub>14</sub> Tyr<sub>15</sub> Asp<sub>16</sub> Ile-  
CCTGAGACTC TCCTGTGCAG CCTCTGGATT CACCTTCAGG AGATATGATA 100  
GGACTCTGAG AGGACACGTC GGAGACCTAA GTGGAAGTCC TCTATACTAT

+1 : - His<sub>1</sub> Trp<sub>2</sub> Val<sub>3</sub> Arg<sub>4</sub> Gln<sub>5</sub> Thr<sub>6</sub> Pro<sub>7</sub> Gly<sub>8</sub> Lys<sub>9</sub> Gly<sub>10</sub> Leu<sub>11</sub> Glu<sub>12</sub> Trp<sub>13</sub> Val<sub>14</sub> Ser<sub>15</sub> Ser-  
TCCACTGGGT CCGCCAGACT CCAGGGAAAGG GCCTGGAGTG GGTCTCATCC 150  
AGGTGACCCA GGCGGTCTGA GGTCCCTTCC CGGACCTCAC CCAGAGTAGG

+1 : - Ile<sub>1</sub> Ser<sub>2</sub> Ser<sub>3</sub> Gly<sub>4</sub> Gly<sub>5</sub> Asn<sub>6</sub> Tyr<sub>7</sub> Ile<sub>8</sub> Asp<sub>9</sub> Tyr<sub>10</sub> Ala<sub>11</sub> Asp<sub>12</sub> Ser<sub>13</sub> Val<sub>14</sub> Lys<sub>15</sub> Gly<sub>16</sub> Arg-  
ATCAGTAGTG GTGGTAATTAA CATAGACTAC GCAGACTCTG TGAAGGGCCG 200  
TAGTCATCAC CACCATTAAAT GTATCTGATG CGTCTGAGAC ACTTCCCAGC

+1 : - Phe<sub>1</sub> Thr<sub>2</sub> Ile<sub>3</sub> Ser<sub>4</sub> Arg<sub>5</sub> Asp<sub>6</sub> Asn<sub>7</sub> Ala<sub>8</sub> Asn<sub>9</sub> Asn<sub>10</sub> Val<sub>11</sub> Val<sub>12</sub> Tyr<sub>13</sub> Leu<sub>14</sub> Gln<sub>15</sub> Met<sub>16</sub> Asn-  
ATTACCATC TCCAGAGACA ACGCCAACAA TGTTGTCTAT CTACAAATGA 250  
TAAGGGTAG AGGTCTCTGT TGCGGTTGTT ACAACAGATA GATGTTTACT

+1 : - Ser<sub>1</sub> Leu<sub>2</sub> Arg<sub>3</sub> Ala<sub>4</sub> Glu<sub>5</sub> Asp<sub>6</sub> Met<sub>7</sub> Ala<sub>8</sub> Val<sub>9</sub> Tyr<sub>10</sub> Phe<sub>11</sub> Cys<sub>12</sub> Ala<sub>13</sub> Arg<sub>14</sub> Asp<sub>15</sub> Gly<sub>16</sub>  
ACAGCCTGAG AGCCGAGGAC ATGGCTGTCT ATTTCTGTGC GAGAGATGGG 300  
TGTGGACTC TCGGCTCCTG TACCGACAGA TAAAGACACG CTCTCTACCC

-1 : - Thr<sub>1</sub> Ile<sub>2</sub> Phe<sub>3</sub> Gly<sub>4</sub> Ser<sub>5</sub> Ala<sub>6</sub> Ala<sub>7</sub> Thr<sub>8</sub> Trp<sub>9</sub> Arg<sub>10</sub> Ala<sub>11</sub> Phe<sub>12</sub> Asp<sub>13</sub> Ile<sub>14</sub> Trp<sub>15</sub> Gly<sub>16</sub> Arg-  
ACGATTTTG GATCGGCGGC GACCTGGCGG GCTTTGATA TCTGGGGCCG 350  
TGCTAAAAAC CTAGCCGCCG CTGGACCGCC CGAAAACATAT AGACCCCCGGC

+1 : - Gly<sub>1</sub> Thr<sub>2</sub> Met<sub>3</sub> Val<sub>4</sub> Thr<sub>5</sub> Val<sub>6</sub> Ser<sub>7</sub> Ser<sub>8</sub>  
GGGGACAATG GTCACCGTGT CGAGT 400  
CCCCCTGTTAC CAGTGGCACCA GCTCA

Figure 9E

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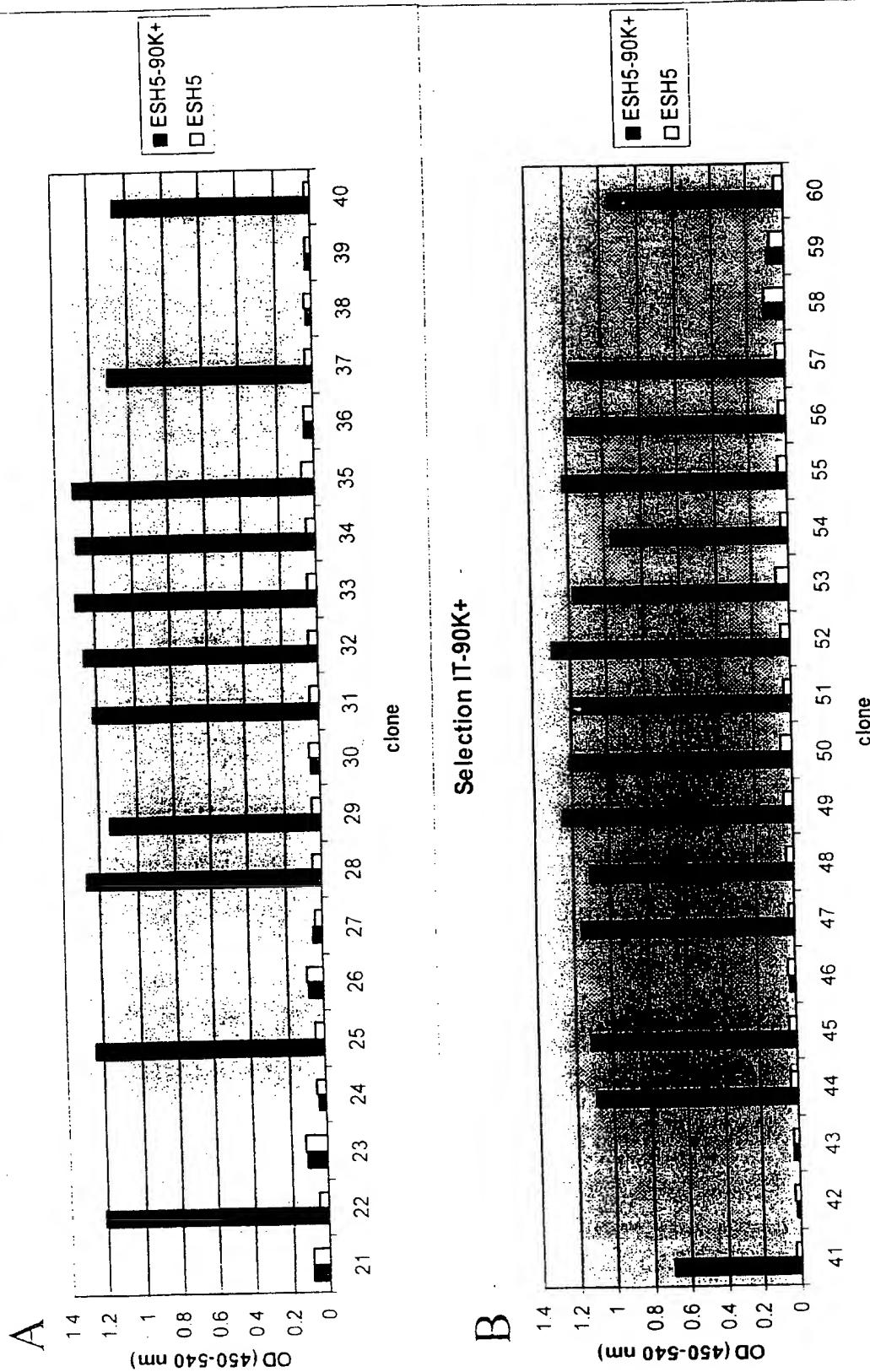


Figure 10

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## Deduced protein sequences of isolated FVIII A2 specific scFv

Heavy chains	FR1	CDR1	FR2	CDR2	FR3	CDR3	FR4
DP-47	EVQLESGGGLVQPQGSRLSCAASGFTS -----V-----D-----V-----	SYAMS WYRQAPKGKGLBEWVS -----A-----NF-----	PIISGSGSTYYADSVKG -----A-----G-RS-T-P-----	RFTISRDNSKNTLYLQMNSLRAEDTAVYYCAK -----V-E-----I-----	RGRGYYKYYGMDV -----V-----	WGGTTVTVSS -----	
DP-10	123456789012345678901234567890 QVQLVQSGAEVKPGFSSVKVSKCAASGFTS -----H-----	12345 678901234567890 12345 SYAIS WYRQAPGQQLEWMG -----D-----L-G-----	012a3456789012345 67890123456789012345 GIIPIGTANTVAQKPGQ RVTITADEBTSSTAYMELSSRSBEDTAVYYCARM -----T-T-----T-T-----	5 6 7 8 9 -----	0 1 1 1 1 -----	567890abccde12 34567890123 -----	34567890123 -----

**Figure 11A**

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+1    rGin, rVal, rGin, rLeu, rVal, rGin, rSer, rGly, rAla, rGlu, rVal, rLys, rLys, rPro, rGly, rSer, rSer-  
CAGGTGCAGC TGGTGCAGTC TGGGGCTGAG GTGAAGAACGC CTGGGTCCCTC 50  
GTCCACGTCG ACCACGTCAG ACCCCGACTC CACTTCTTCG GACCCAGGAG

+1    ..rVal, rLys, rVal, rSer, rCys, rLys, rAla, rSer, rGly, rGly, rThr, rPhe, rSer, rSer, rHis, rAla, rIle-  
GGTGAAGGTC TCCTGCAAGG CTTCTGGAGG CACCTTCAGC AGTCATGCTA 100  
CCACTTCCAG AGGACGTTCC GAAGACCTCC GTGGAAQTCG TCAGTACGAT

+1    ...rSer, rTrp, rVal, rArg, rGin, rAla, rPro, rGly, rGin, rGly, rLeu, rGlu, rTrp, rMet, rGly, rAsp-  
TCAGCTGGGT GCGACAGGCC CCTGGACAAG GGCTTGAGTG GATGGGAGAC 150  
AGTCGACCCA CGCTGTCCGG GGACCTGTTCC CCGAACTCAC CTACCCTCTG

+1    ...rIle, rIle, rPro, rIle, rLeu, rGly, rThr, rGly, rAsn, rTyr, rAla, rGln, rLys, rPhe, rGln, rGly, rArg-  
ATCATCCCTA TCCTTGGTAC AGGAAACTAC GCACAGAAAGT TCCAGGGCAG 200  
TAGTAGGGAT AGGAACCATG TCCTTGATG CGTGTCTTCA AGGTCCCCGTC

+1    ...rVal, rThr, rIle, rThr, rAla, rAsp, rGlu, rSer, rThr, rSer, rThr, rAla, rTyr, rMet, rGlu, rLeu, rSer-  
AGTCACGATT ACCGCGGACG AGTCCACGAG CACAGCCTAC ATGGAGCTGA 250  
TCAGTGCTAA TGGCGCCTGC TCAGGTGCTC GTGTCGGATG TACCTCGACT

+1    ...rThr, rLeu, rThr, rSer, rGlu, rAsp, rThr, rAla, rVal, rTyr, rTyr, rCys, rGlu, rLeu, rAsp, rTrp-  
GCACCCCTGAC ATCTGAGGAC ACGGCCGTGT ATTACTGTGA ACTTGACTGG 300  
CGTGGGACTG TAGACTCCTG TGCCGGCACA TAATGACACT TGAACGTGACC

-1    ...rPhe, rTyr, rIle, rTrp, rGly, rGln, rGly, rThr, rMet, rVal, rThr, rVal, rSer, rSer-  
TTCTATATCT GGGGCCAAGG GACAATGGTC ACCGTGTCGA GT 350  
AAGATATAGA CCCCGGTTCC CTGTTACAG TGGCACAGCT CA

Figure 11B

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+1    [-Glu-] Val [-Gln-] [-Leu-] Val [-Glu-] Ser [-Gly-] Gly [-Asp-] [-Leu-] Val [-Gln-] Pro [-Gly-] Gly [-Ser-]  
 GAGGTGCAGC TGGTGGAGTC TGGGGGAGAC TTGGTACAGC CTGGGGGGTC 50  
 CTCCACGTAG ACCACCTCAG ACCCCCTCTG AACCATGTCG GACCCCCCAG

+1    [-Leu-] Arg [-Leu-] Ser [-Cys-] Ala [-Ala-] Ser [-Gly-] Phe [-Thr-] Phe [-Ser-] Asn [-Phe-] Ala [-Met-]  
 CCTGAGACTC TCCTGTGCAG CCTCTGGATT CACCTTAGC AACTTGCCA 100  
 GGACTCTGAG AGGACACGTC GGAGACCTAA GTGGAAATCG TTGAAACGGT

+1    [-Ser-] Trp [-Val-] Arg [-Gln-] Ala [-Pro-] Gly [-Lys-] Gly [-Leu-] Glu [-Trp-] Val [-Ala-] Ala [-Ala-]  
 TGAGCTGGGT CCGCCAGGCT CCCGGGAAGG GGCTGGAGTG GGTCGCGGCT 150  
 ACTCGACCCA GGCGGTCCGA GGGCCCTTCC CCGACCTCAC CCAGCGCCGA

+1    [-Ile-] Gly [-Gly-] Arg [-Ser-] Gly [-Thr-] Thr [-Phe-] Tyr [-Ala-] Asp [-Ser-] Val [-Lys-] Gly [-Arg-]  
 ATTGGCGGTA GAAGTGGTAC CACATTCTAC GCGGACTCCG TGAAGGGCCG 200  
 TAACCGCCAT CTTCACCATG GTGTAAGATG CGCCTGAGGC ACTTCCCCGGC

+1    [-Phe-] Thr [-Ile-] Ser [-Arg-] Asp [-Asn-] Ser [-Lys-] Asn [-Thr-] Val [-Tyr-] Leu [-Glu-] Met [-Asn-]  
 GTTCACCATC TCCAGAGACA ATTCCAAGAA CACGGTCTAT CTGGAAATGA 250  
 CAAGTGGTAG AGGTCTCTGT TAAGGTTCTT GTGCCAGATA GACCTTACT

+1    [-Ser-] Leu [-Arg-] Ala [-Glu-] Asp [-Thr-] Ala [-Ile-] Tyr [-Tyr-] Cys [-Ala-] Lys [-Arg-] Gly [-]  
 ACAGTCTGAG AGCCGAGGAC ACAGCCATTT ATTACTGTGC GAAAAGAGGG 300  
 TGTCAGACTC TCGGCTCCTG TGTGGTAAA TAATGACACG CTTTCTCCC

+1    [-Arg-] Gly [-Gly-] Tyr [-Lys-] Tyr [-Tyr-] Gly [-Met-] Asp [-Val-] Trp [-Gly-] Gln [-Gly-] Thr [-Thr-]  
 CGCGGGGGGT ATAAGTATTA TGGGATGGAC GTCTGGGGCC AGGGGACAC 350  
 GCGCCCCCAGA TATTCTACCTG CAGACCCCGG TCCCCCTGGTG

+1    .. [-Val-] Thr [-Val-] Ser [-Ser-]  
 GGTCACCGTG TCGAGT  
 CCAGTGGCAC AGCTCA

400

Figure 11C